10/763,935 December 27, 2007

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```
=> d que 16
            19 SEA FILE=REGISTRY ABB=ON PLU=ON PP.S.D..F...R.....QE.
L5
               ..R..../SQSP
L6
            17 SEA FILE=REGISTRY ABB=ON PLU=ON L5 AND (SOL=38 OR SOL=39)
=> d 16 rn cn sql kwic lc nte tot
    ANSWER 1 OF 17 REGISTRY COPYRIGHT 2007 ACS on STN
L6
    935739-49-4 REGISTRY
RN
    L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-
     L-\alpha-aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-
     L-leucyl-L-arginyl-L-\alpha-glutamyl-L-valyl-L-leucyl-L-\alpha-glutamyl-
     L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L-\alpha-glutamyl-L-glutaminyl-N-
     methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L-\alpha-glutamyl-L-
     histidyl-L-seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L-\alpha-
     qlutamyl-N-methyl-L-leucyl-, (28\rightarrow31)-lactam (CA INDEX NAME)
SQL 38
SOL 38
         1 PPISLDLTFH LLREVLEXAR AEQLAQQEHS KRKLXELI
          ______ ____
HITS AT: 1-38
**RELATED SEQUENCES AVAILABLE WITH SEQLINK**
    STN Files: CA, CAPLUS
```

type		 location	description
bridge	Glu-28	- Lys-31	lactam
uncommon	Nle-18	_	_
uncommon	Nle-35	_	_

NTE modified (modifications unspecified)

Phe-9 D ANSWER 2 OF 17 REGISTRY COPYRIGHT 2007 ACS on STN 935739-47-2 REGISTRY RN CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl- $L-\alpha$ -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl- $L-leucyl-L-arginyl-L-\alpha-glutamyl-L-valyl-L-leucyl-L-\alpha-glutamyl-$ L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L-qlutaminyl-Nmethyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-Lhistidyl-L-seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α glutamyl-N-methyl-L-leucyl- (CA INDEX NAME) SQL 38 38 SQL SEQ 1 PPISLDLTFH LLREVLEXAR AEOLAQOEHS KRKLXELI HITS AT: 1-38 **RELATED SEQUENCES AVAILABLE WITH SEQLINK** LC STN Files: CA, CAPLUS NTE modified (modifications unspecified) ----- location ----description ______ Nle-18 uncommon Nle-35 uncommon Phe-9 – D L6 ANSWER 3 OF 17 REGISTRY COPYRIGHT 2007 ACS on STN RN 935739-46-1 REGISTRY CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl- $L-\alpha$ -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl- $L-leucyl-L-arginyl-L-\alpha-glutamyl-L-valyl-L-leucyl-L-\alpha-glutamyl L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L-\alpha-qlutamyl-L-qlutaminyl-L$ $leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L-\alpha-glutamyl-L-histidyl-L$ $seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L-\alpha-glutamyl-L$ isoleucyl-, $(28\rightarrow31)$ -lactam (CA INDEX NAME) OTHER NAMES: CMStressin1-A SQL 38 SQL 38 SEQ 1 PPISLDLTFH LLREVLEXAR AEQLAQQEHS KRKLXEII _____ ___ ___ HITS AT: 1-38 **RELATED SEQUENCES AVAILABLE WITH SEQLINK** LC STN Files: CA, CAPLUS, PROUSDDR NTE modified _____ ----- location ----description terminal mod. Pro-1 - N-acetyl terminal mod. Ile-38 - C-terminal amide bridge Glu-28 - Lys-31 lactam uncommon Nle-18 - -

10/763,935 December 27, 2007

Nle-35 uncommon Phe-9 D stereo ANSWER 4 OF 17 REGISTRY COPYRIGHT 2007 ACS on STN 935739-45-0 REGISTRY RN CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl- $L-\alpha$ -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl- $L-leucyl-L-arginyl-L-\alpha-glutamyl-L-valyl-L-leucyl-L-\alpha-glutamyl-ucyl-L-\alpha-glutamyl-ucyl-L-arginyl$ $L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L-\alpha-glutamyl-L-glutaminyl-L$ $leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L-\alpha-glutamyl-L-histidyl-L$ $seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L-\alpha-glutamyl-L$ isoleucyl- (CA INDEX NAME) SQL 38 SQL 38 SEO 1 PPISLDLTFH LLREVLEXAR AEQLAQQEHS KRKLXEII ______ _____ HITS AT: 1-38 **RELATED SEQUENCES AVAILABLE WITH SEQLINK** LC STN Files: CA, CAPLUS NTE modified ______ ----- location ----- description C-terminal amide Phe-9 stereo L6 ANSWER 5 OF 17 REGISTRY COPYRIGHT 2007 ACS on STN 496031-25-5 REGISTRY RN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl- $L-\alpha$ -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl- $L-leucyl-L-arginyl-L-\alpha-glutamyl-L-valyl-L-leucyl-L-\alpha-glutamyl-$ L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L-qlutaminyl-L $leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L-\alpha-glutamyl-L-histidyl-L$ $seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L-\alpha-glutamyl-L$ isoleucyl- (9CI) (CA INDEX NAME) SOL 38 3.8 SQL SEO 1 PPISLDLTFH LLREVLEXAR AEQLAQQEHS KRKLXEII HITS AT: 1-38 **RELATED SEQUENCES AVAILABLE WITH SEQLINK** LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL NTE modified ----- location ----- description terminal mod. Pro-1 terminal mod. Ile-38 uncommon Nle-18 N-acetyl C-terminal amide

Nle-35 uncommon Phe-9 D stereo ANSWER 6 OF 17 REGISTRY COPYRIGHT 2007 ACS on STN RN 496031-24-4 REGISTRY CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl- $L-\alpha$ -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl- $L-leucyl-L-arginyl-L-\alpha-glutamyl-L-valyl-L-leucyl-L-\alpha-glutamyl-ucyl-L-\alpha-glutamyl-ucyl-L-arginyl$ $L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L-\alpha-glutamyl-L-glutaminyl-2$ methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-Lhistidyl-D-alanyl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α glutamyl-2-methyl-L-leucyl- (9CI) (CA INDEX NAME) SQL 38 SQL 38 SEO 1 PPISLDLTFH LLREVLEXAR AEQLAQQEHA KRKLXELI ______ _____ HITS AT: 1-38 **RELATED SEQUENCES AVAILABLE WITH SEQLINK** LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL NTE modified (modifications unspecified) _____ ----- location ----- description uncommon Nle-18 - uncommon Nle-35 -Phe-9 stereo Ala-30 D stereo L6 ANSWER 7 OF 17 REGISTRY COPYRIGHT 2007 ACS on STN RN 496031-23-3 REGISTRY CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl- $L-\alpha$ -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl- $L-leucyl-L-arginyl-L-\alpha-glutamyl-L-valyl-L-leucyl-L-\alpha-glutamyl L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L-\alpha-qlutamyl-L-qlutaminyl-2$ $methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L-\alpha-glutamyl-L$ histidyl-D-alanyl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α glutamy1-2-methy1-L-leucy1-, $(28\rightarrow31)$ -lactam (9CI) (CA INDEX NAME) SQL 38 SQL 38 SEQ 1 PPISLDLTFH LLREVLEXAR AEQLAQQEHA KRKLXELI _____ ___ ___ HITS AT: 1-38 **RELATED SEQUENCES AVAILABLE WITH SEQLINK** LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL NTE modified (modifications unspecified) ______ ----- location ----description bridge Glu-28 - Lys-31 lactam uncommon Nle-18 - - - - - stereo Phe-9 - D

D

Phe-9

stereo

stereo Ala-30 - D

L6 ANSWER 8 OF 17 REGISTRY COPYRIGHT 2007 ACS on STN

RN 496031-22-2 REGISTRY

CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L- α -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-leucyl-L-arginyl-L- α -glutamyl-L-valyl-L-leucyl-L- α -glutamyl-L-leucyl-L-alanyl-L-alanyl-L-alanyl-L-glutaminyl-2-methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-L-histidyl-D-seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α -glutamyl-2-methyl-L-leucyl- (9CI) (CA INDEX NAME)

SQL 38 SQL 38

SEQ 1 PPISLDLTFH LLREVLEXAR AEQLAQQEHS KRKLXELI

HITS AT: 1-38

RELATED SEQUENCES AVAILABLE WITH SEQLINK

LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

NTE modified (modifications unspecified)

type ----- location ----- description

uncommon Nle-18 - uncommon Nle-35 - stereo Phe-9 - D
stereo Ser-30 - D

L6 ANSWER 9 OF 17 REGISTRY COPYRIGHT 2007 ACS on STN

RN 496031-21-1 REGISTRY

CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L- α -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-leucyl-L-arginyl-L- α -glutamyl-L-leucyl-L- α -glutamyl-L-alanyl-L-alanyl-L-alanyl-L-glutaminyl-2-methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-L-histidyl-D-seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α -glutamyl-2-methyl-L-leucyl-, (28 \rightarrow 31)-lactam (9CI) (CA INDEX NAME)

SQL 38

SEQ 1 PPISLDLTFH LLREVLEXAR AEQLAQQEHS KRKLXELI

HITS AT: 1-38

RELATED SEQUENCES AVAILABLE WITH SEQLINK
LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL
NTE modified (modifications unspecified)

type	lo	cation	description	
bridge	Glu-28	- Lys-31	lactam	
uncommon	Nle-18	_	_	
uncommon	Nle-35	_	_	
stereo	Phe-9	_	D	
stereo	Ser-30	_	D	

```
L6
    ANSWER 10 OF 17 REGISTRY COPYRIGHT 2007 ACS on STN
RN
    496031-20-0 REGISTRY
CN
    L-Isoleucinamide, 1-acetyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-
    L-\alpha-aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-
    L-leucyl-L-arginyl-L-\alpha-glutamyl-L-valyl-L-leucyl-L-\alpha-glutamyl-
    L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L-\alpha-glutamyl-L-glutaminyl-2-
    methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L-qlutaminyl-L-
    histidyl-2-methylalanyl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L-
    \alpha-glutamyl-2-methyl-L-leucyl- (9CI) (CA INDEX NAME)
SQL
SQL
    38
SEQ
        1 PPISLDLTFH LLREVLEXAR AEQLAQQEHX KRKLXELI
         _____ ___ ___
HITS AT: 1-38
**RELATED SEOUENCES AVAILABLE WITH SEOLINK**
LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL
NTE modified (modifications unspecified)
______
              ----- location -----
                                      description
______
uncommon Nle-18
uncommon Aib-30
uncommon Nle-35
stereo Phe-9
                            – D
L6 ANSWER 11 OF 17 REGISTRY COPYRIGHT 2007 ACS on STN
RN 496031-19-7 REGISTRY
CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-
    L-\alpha-aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-
    L-leucyl-L-arginyl-L-\alpha-glutamyl-L-valyl-L-leucyl-L-\alpha-glutamyl-
    L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L-\alpha-glutamyl-L-glutaminyl-2-
    methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L-\alpha-glutamyl-L-
    histidyl-2-methylalanyl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L-
    \alpha-glutamyl-2-methyl-L-leucyl-, (28\rightarrow31)-lactam (9CI) (CA
    INDEX NAME)
SOL
    38
SOL
    38
SEO
        1 PPISLDLTFH LLREVLEXAR AEQLAQQEHX KRKLXELI
         _____ ____
HITS AT:
         1 - 38
**RELATED SEQUENCES AVAILABLE WITH SEQLINK**
    STN Files: CA, CAPLUS, TOXCENTER, USPATFULL
NTE modified (modifications unspecified)
______
              ----- location ----- description
tvpe
```

```
L6
        ANSWER 12 OF 17 REGISTRY COPYRIGHT 2007 ACS on STN
RN
        496031-18-6 REGISTRY
CN
        L-Isoleucinamide, 1-acetyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-
        L-\alpha-aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-
         L-leucyl-L-arginyl-L-\alpha-glutamyl-L-valyl-L-leucyl-L-\alpha-glutamyl-
         L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L-\alpha-glutamyl-L-glutaminyl-2-
         methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L-\alpha-glutamyl-L-
         histidyl-L-seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L-\alpha-
         glutamyl-2-methyl-L-leucyl- (9CI) (CA INDEX NAME)
SQL
        3.8
SQL
       38
SEQ
                1 PPISLDLTFH LLREVLEXAR AEQLAQQEHS KRKLXELI
                   _____ ___ ___
                1-38
HITS AT:
**RELATED SEOUENCES AVAILABLE WITH SEOLINK**
      STN Files: CA, CAPLUS, TOXCENTER, USPATFULL
NTE modified (modifications unspecified)
______
                             ----- location -----
                                                                                   description
______
uncommon Nle-18 uncommon Nle-35
stereo Phe-9
                                                                          D
1.6
        ANSWER 13 OF 17 REGISTRY COPYRIGHT 2007 ACS on STN
        496031-17-5 REGISTRY
RN
CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-
        L-\alpha-aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-
         L-leucyl-L-arginyl-L-\alpha-glutamyl-L-valyl-L-leucyl-L-\alpha-glutamyl-ucyl-L-\alpha-glutamyl-ucyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl
         L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L-\alpha-glutamyl-L-glutaminyl-2-
         methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L-a-a-glutamyl-L-
         histidyl-L-seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L-\alpha-
         qlutamyl-2-methyl-L-leucyl-, (28\rightarrow31)-lactam (9CI) (CA INDEX NAME)
SOL
        38
SOL
       38
SEQ
                1 PPISLDLTFH LLREVLEXAR AEQLAQQEHS KRKLXELI
                    _____ ____
HITS AT:
                1-38
**RELATED SEQUENCES AVAILABLE WITH SEQLINK**
        STN Files: CA, CAPLUS, TOXCENTER, USPATFULL
NTE modified (modifications unspecified)
_____
                              ----- location ----- description
 tvpe
______
                      Glu-28 - Lys-31 lactam
Nle-18 - -
uncommon
                           Nle-35
uncommon
                      Phe-9
                                                                  D
stereo
```

```
10/763,935
RN
                496031-16-4 REGISTRY
CN
               L-Leucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L-
                \alpha-aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-
                leucyl-L-arginyl-L-\alpha-glutamyl-L-valyl-L-leucyl-L-\alpha-glutamyl-L-
                norleucyl-L-alanyl-L-arginyl-L-alanyl-L-\alpha-glutamyl-L-glutaminyl-2-
                methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L-q-glutamyl-L-
                histidyl-2-methylalanyl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L-
                \alpha-glutamyl-L-isoleucyl-2-methyl-, (28\rightarrow31)-lactam (9CI) (CA
                INDEX NAME)
              38
SQL
SQL 38
SEQ
                            1 PPISLDLTFH LLREVLEXAR AEQLAQQEHX KRKLXEIL
                                   ______ ____
                            1-38
LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL
NTE modified (modifications unspecified)
 ______
                                                  ----- location -----
                                                                                                                                                  description
 ______

      bridge
      Glu-28
      - Lys-31
      lactam

      uncommon
      Nle-18
      -
      -

      uncommon
      Aib-30
      -
      -

      uncommon
      Nle-35
      -
      -

ANSWER 15 OF 17 REGISTRY COPYRIGHT 2007 ACS on STN
L6
RN
               496031-15-3 REGISTRY
CN
               L-Leucinamide, 1-acetyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L-
               \alpha-aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-
                \texttt{leucyl-L-arginyl-L-}\alpha - \texttt{glutamyl-L-valyl-L-leucyl-L-}\alpha - \texttt{glutamyl-L-valyl-L-leucyl-L-}\alpha - \texttt{glutamyl-L-}\alpha - \texttt{gl
                norleucyl-L-alanyl-L-arginyl-L-alanyl-L-\alpha-glutamyl-L-glutaminyl-2-
                methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L-\alpha-glutamyl-L-
                histidyl-L-seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L-\alpha-
                qlutamyl-L-isoleucyl-2-methyl-, (28\rightarrow 31)-lactam (9CI) (CA INDEX
               NAME)
SOL
              38
              38
```

SQL

SEO 1 PPISLDLTFH LLREVLEXAR AEQLAQQEHS KRKLXEIL ______ ____

HITS AT: 1-38

LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

NTE modified (modifications unspecified)

______ ----- location ----description ______ bridge Glu-28 uncommon Nle-18 uncommon Nle-35 stereo Phe-9 Glu-28 - Lys-31 Nle-18 -Nle-35 lactam D

- ANSWER 16 OF 17 REGISTRY COPYRIGHT 2007 ACS on STN L6
- 496031-14-2 REGISTRY RN
- L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl- $L-\alpha$ -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-

```
L-leucyl-L-arginyl-L-\alpha-glutamyl-L-valyl-L-leucyl-L-\alpha-glutamyl-ucyl-L-\alpha-glutamyl-ucyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl-L-arginyl
          L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L-\alpha-glutamyl-L-glutaminyl-L-
          leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L-\alpha-glutamyl-L-histidyl-L-
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NTE modified
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terminal mod. Pro-1 - terminal mod. Ile-38 - bridge Glu-28 - Lys-31 uncommon Nle-18 - uncommon Nle-35 -
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                              Phe-9
stereo
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      ANSWER 17 OF 17 REGISTRY COPYRIGHT 2007 ACS on STN
L6
        462692-83-7 REGISTRY
RN
CN Peptide, (Pro-Pro-Leu-Ser-Ile-Asp-Leu-Thr-Phe-Xaa-Leu-Leu-Arg-Asn-Met-Met-
         Gln-Arg-Ala-Glu-Met-Glu-Lys-Leu-Arg-Glu-Gln-Glu-Lys-Ile-Asn-Arg-Glu-Ile-
         Leu-Glu-Gln-Val) (9CI) (CA INDEX NAME)
OTHER NAMES:
        11: PN: WO02074326 SEQID: 12 unclaimed protein
SQL 38
SOL 38
                 1 PPLSIDLTFX LLRNMMQRAE MEKLREQEKI NREILEQV
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L5
19 SEA FILE=REGISTRY ABB=ON PLU=ON PP.S.D..F...R......QE.
..R..../SQSP
L6
17 SEA FILE=REGISTRY ABB=ON PLU=ON L5 AND (SQL=38 OR SQL=39)
L8
3 SEA FILE=CAPLUS ABB=ON PLU=ON L6
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AUTHOR(S):

CORPORATE SOURCE:

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L8 ANSWER 1 OF 3 CAPLUS COPYRIGHT 2007 ACS on STN ACCESSION NUMBER: 2007:247482 CAPLUS <u>Full-text</u>
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DOCUMENT NUMBER: 146:474752

TITLE: Stressin1-A, a Potent Corticotropin Releasing Factor

Receptor 1 (CRF1)-Selective Peptide Agonist Rivier, Jean; Gulyas, Jozsef; Kunitake, Koichi; DiGruccio, Michael; Cantle, Jeffrey P.; Perrin,

Marilyn H.; Donaldson, Cindy; Vaughan, Joan; Million, Mulugeta; Gourcerol, Guillaume; Adelson, David W.;

Rivier, Catherine; Tache, Yvette; Vale, Wylie The Clayton Foundation Laboratories for Peptide

Biology, The Salk Institute for Biological Studies, La

Jolla, CA, 92037, USA

SOURCE: Journal of Medicinal Chemistry (2007), 50(7),

1668-1674

CODEN: JMCMAR; ISSN: 0022-2623

PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal LANGUAGE: English

The potencies and selectivity of peptide CRF antagonists is increased through structural constraints, suggesting that the resulting ligands assume distinct conformations when interacting with CRF1 and CRF2 receptors. To develop selective CRF receptor agonists, we have scanned the sequence -Gln-Ala-His-Ser-Asn-Arg- (residues 30-35 of [DPhe12,Nle21,38]Ac-hCRF4-41) with an i-(i+3) bridge consisting of the Glui-Xaa-Xbb-Lysi+3 scaffold, where residues i = 30, 31, and 32. When i = 31, stressin1-A, a potent CRF1 receptor-selective agonist was generated. In vitro, stressin1-A was equipotent to h/rCRF to release ACTH. Astressin1-A showed a low nanomolar affinity for CRF1 receptor (Ki = 1.7 nM) and greater than 100-fold selectivity vs. CRF2 receptor (Ki = 222 nM). Stressin1-A released slightly less ACTH than oCRF in adult adrenal-intact male rats, with increased duration of action. Stressin1-A, injected i.p. in rats, induced fecal pellet output (a CRF1 receptor-mediated response) and did not influence gastric emptying and blood pressure (CRF2 receptor-mediated responses).

IT 935739-45-0P 935739-46-1P, Stressin1-A 935739-47-2P 935739-49-4P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); BIOL

(Biological study); PREP (Preparation)

(Stressin1-A as CRF1-selective peptide agonist)

REFERENCE COUNT: 34 THERE ARE 34 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 2 OF 3 CAPLUS COPYRIGHT 2007 ACS on STN ACCESSION NUMBER: 2003:117793 CAPLUS Full-text

DOCUMENT NUMBER: 138:153832

Preparation of corticotropin-releasing factor (CRF) TITLE:

analogs as CRF receptor type 1 (CRFR1) selective

ligands

Rivier, Jean E. F.; Vale, Wylie W., Jr.; Perrin, INVENTOR(S):

Marilyn H.; Guylas, Jozsef

PATENT ASSIGNEE(S): The Salk Institute for Biological Studies, USA

SOURCE: PCT Int. Appl., 43 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PA	PATENT NO.						KIND DATE				ICAT	ION 1		DATE				
				A2 20030213 A3 20070920			•	 WO 2	002-	US24	238	20020730						
	W:	ΑE,	AG,	AL,	AM,	ΑT,	ΑU,	AZ,	BA,	BB,	BG,	BR,	BY,	BZ,	CA,	CH,	CN,	
		CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,	
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		LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NO,	NΖ,	OM,	PH,	
		PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,	ΤJ,	TM,	TN,	TR,	TT,	TZ,	
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		PT,	SE,	SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	
		ΝE,	SN,	TD,	ΤG,	AP,	EA,	AM,	AZ,	BY,	KG,	KΖ,	MD,	RU,	ТJ,	TM,	EP,	OA
CA	2455	223			A1 20030213 CA						CA 2002-2455223					20020730		
AU	2002	3557	42		A1 20030217					AU 2002-355742					20020730			
JP	2005	5104	58		Τ		2005	0421	1	JP 2	003-	5170	15	20020730				
EP	1572	679			A2		2005	0914	EP 2002-752639					20020730				
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US	US 2004204564						2004	1014	US 2004-763935					20040122				
PRIORIT	Y APP	LN.	INFO	.:						US 2	001-	3095	04P		P 20010801			
							WO 2002-US24238						238	W 20020730				

MARPAT 138:153832 OTHER SOURCE(S):

Corticotropin-releasing factor (CRF) peptides Y1-Pro-Pro-R6-Ser-R8-Asp-R10-R11-D-Phe-R13-R14-R15-Arg-R17-R18-R19-R20-R21-R22-R23-R24-R25-R26-R27-R28-R29-Gln-Glu-R32-R33-R34-Arg-R36-R37-R38-R39-R40-R41-NH2 (Y1 is acyl having < 15 carbon atoms or radioiodinated tyrosine; the R groups represent various amino acid residues which are defined) or their nontoxic salts are claimed for selective binding to CRFR1. Thus, cyclo(31-34)(Ac-Pro4,D-Phe12,Nle21,38,Glu31,Lys34)-r/hCRF(4-41) was prepared by the solid-phase method and shown to bind hCRFR1 with high affinity and significantly lowered blood pressure when administered peripherally.

ΙT 496031-18-6P 496031-20-0P 496031-22-2P 496031-24-4P 496031-25-5P

> RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(preparation of corticotropin-releasing factor (CRF) analogs as CRF

receptor

type 1 (CRFR1) selective ligands)
IT 496031-14-2P 496031-15-3P 496031-16-4P
496031-17-5P 496031-19-7P 496031-21-1P
496031-23-3P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

type 1 (CRFR1) selective ligands)

L8 ANSWER 3 OF 3 CAPLUS COPYRIGHT 2007 ACS on STN ACCESSION NUMBER: 2002:736126 CAPLUS Full-text

DOCUMENT NUMBER: 137:257953

TITLE: A new human urocortin identified by sequence homology

acting as an agonist for type II corticotropin

releasing factor receptors

INVENTOR(S): Vale, Wylie W., Jr.; Rivier, Jean E.; Kunitake, Koichi

S.; Lewis, Kathy A.; Perrin, Marilyn H.; Gulyas,

Jozsef

PATENT ASSIGNEE(S): Research Development Foundation, USA

SOURCE: PCT Int. Appl., 81 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

										APPLICATION NO.										
WO	2002	0743	26		A2		2002		WO 2002-US9115											
WO	2002								BA, BB, BG, BR, BY, BZ											
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						CA 2002-2425902														
AU	2002306853			A1	A1 20021003			AU 2002-306853							20020315					
US	2003	2003036507					20030220 US 2002-99766					20020315								
US	6812	210			В2		2004	1102												
EP	1368	051			A2		2003	1210	EP 2002-753685							20020315				
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NZ	5244	77			Α		20041126 NZ 2002-524477													
JP	2005						20050224 JP 2002-573033							20020315						
	2305	109			C2		2007	0827		RU	20	03-	1049	77		2	20020	315		
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	6953				В2		2005	1011												
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AU	AU 2006225288 A			A1									88			20061	006			
RIORIT	Y APP	LN.	INFO	.:						US	20	01-	2760	69P		P 2	20010	315		
										US	20	01-	2949	14P		P 2	20010	531		
										US	20	02-	9976	6		A3 2	20020	315		

A1 20040203

WO 2002-US9115 W 20020315 US 2004-771224

AΒ A search of the public human genome database identified a human EST (GenBank AW293249) highly similar pufferfish urocortin sequences. The full length sequence was amplified from human genomic DNA and sequenced. Sequence homol. comparisons of the novel sequence with human urocortin I and urocortin II revealed that the sequence encoded a novel human urocortin, which was designated urocortin III (UcnIII). While urocortin III does not have high affinity for either of the corticotropin releasing factor receptors CRF-R1 or CRF-R2, the affinity for CRF-R2 is greater than the affinity for CRF-R1. Urocortin III is capable stimulating cAMP production in cells expressing CRF-Thus, the affinity is high enough that urocortin III could act as a native agonist of CRF-R2 and so could be the lead compound for the development of therapeutics acting on the receptor. N-terminal deletion derivs. of urocortin III were effective antagonists. However, it is also likely that urocortin III is a stronger agonist of a yet to be identified receptor. 462692-83-7

RL: PRP (Properties)

(unclaimed protein sequence; new human urocortin identified by sequence homol. acting as an agonist for type II corticotropin releasing factor receptors)

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ΙT

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12 SEA ABB=ON PLU=ON L2 AND (SQL=38 OR SQL=39) L3

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L519 SEA ABB=ON PLU=ON PP.S.D..F...R.....QE...R..../SQSP

L6 17 SEA ABB=ON PLU=ON L5 AND (SQL=38 OR SQL=39)

12 SEA ABB=ON PLU=ON L6 AND L3 L7

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FILE 'REGISTRY' ENTERED AT 13:05:28 ON 27 DEC 2007 5 SEA ABB=ON PLU=ON L6 NOT L7 L9

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10/763,935 December 27, 2007

D L6 RN CN SQL KWIC LC NTE TOT

FILE 'CAPLUS' ENTERED AT 13:11:01 ON 27 DEC 2007 D QUE L8

D L8 IBIB ABS HITRN TOT